## In the Claims:

This listing of claims will replace all prior versions, and listings, of claims in the application. New material is indicated by an <u>underline</u>, deleted material is indicated by a <u>strikethrough</u>.

## Listing of claims:

 (Currently Amended) A method of preparing stable, purely synthetic, self-assembling, controlled release, polyethylene oxide (PEO)-based polymersome vesicles having a semipermeable, thin-walled, amphiphilic, PEO-based block copolymer encapsulating membrane and at least one active agent encapsulated therein to form at least one encapsulant, the method comprising:

determining the appropriate blend ratio (mol %) of a hydrolysable PEO-based block copolymer of having at least one hydrophilie polyester component, and at least one more hydrophobie inert PEO-based block copolymer emponent having at least one hydrophobic component, to produce amphiphilic PEO-based polymersomes having a desired controlled release rate of the encapsulated encapsulant based upon the blend ratio;

selecting the at least one hydrolytically degradable, hydrophobic inert PEO-based block copolymer to effect controlled polyester chain hydrolysis in the membrane, such that when combined with hydrophilie the hydrolysable PEO-based block copolymer, the PEO volume fraction ( $f_{EO}$ ) and chain chemistry control encapsulant release kinetics from the copolymer vesicles, and further control polymersome carrier membrane destabilization; and

blending in aqueous solution the at least one hydrophilie hydrolysable PEO-based block copolymer together with the at least one inert, hydrophobie PEO-based block copolymer to effect self-assembly without secondary chemical processing of the amphiphilic PEO-based polymersomes, without the use of a co-solvent, having the desired controlled release rate of the at least one encapsulant contained therein when the encapsulant is released by hydrolysis-driven membrane poration.

(Currently Amended) The method of claim 1, wherein the polyethylene oxide component
of both the hydrolysable PEO-based block copolymer and the inert PEO-based block copolymer

is polyethylene glycol (PEG).

## (Cancelled)

- (Currently Amended) The method of claim 3 2, wherein the hydrolytically degradable
  polyester component of the hydrolysable PEO-based block copolymer comprises a polyester of
  polylactic acid (PLA) or a polycaprolactone (PCL).
- (Currently Amended) The method of claim 1, wherein the <u>hydrophobic component of the</u>
  at least one inert, non hydrophilie PEO-based block copplymer comprises polybutadiene.
- 6. (Currently Amended) The method of claim 1, wherein the rate of controlled release of the encapsulant upon subsequent hydration of the polymersome is a linear function of an initial (mol %) of the at least one hydrolytically degradable inert PEO- based block copolymer in the blend ratio.
- (Currently Amended) The method of claim 6, wherein increasing the block f<sub>EO</sub> increases
  the rate of transformation into a detergent-like moiety, thereby accelerating destabilization of
  bilayer morphology of the polymersome membrane and encapsulant release.
- (Currently Amended) The method of claim 1, further comprising selecting the at-least-one
  polyester component of the hydrolysable PEO-based block copolymer for biocompatibility.
- (Original) The method of claim 1, wherein the at least one encapsulant is an amphiphilic or lipophilic composition.
- (Previously Presented) The method of claim 1, wherein the at least one encapsulant ranges in molecular weight from 10<sup>2</sup> Da to 10<sup>5</sup> Da.
- 11. (Currently Amended) The method of claim 1, wherein increasing the molecular weight of the at least one encapsulant decelerates the rate of release from the polymersome carrier.

PHIP/744092.1

- 12. (Currently Amended) The method of claim 9, wherein the at least one encapsulant is a hydrophilic encapsulant encapsulated in the lumen of the polymersome, or the at least one encapsulant is a hydrophilic encapsulant encapsulated by intercalation into the polymersome membrane, or the encapsulant is more than more one encapsulant selected from one or more hydrophilic encapsulants, or a combination thereof.
- 13. (Currently Amended) The method of claim 12, wherein at least one hydrophilic encapsulant is selected from the group consisting of carbohydrates, sucrose; marker-tagged dextrans, fluorescent dextrans from 1 kD up to 200 kD; therapeutic compositions, doxorubicin or amphoterican B; dyes; indicators; protein or protein fragments, catalase; ammonium sulfate; salts; and gene, of gene fragments, or oligonucleotides.
- 14. (Previously Presented) The method of claim 12, wherein at least one hydrophobic encapsulant is selected from the group consisting of PKH fluorescent dyes; therapeutic compositions, taxol and anthracyclin; monosialoganglioside; fluorinated lipids; fluorescein-taxol; and fluorescent-dye modified copolymers.
- (Currently Amended) The method of claim 12, wherein the at least one encapsulant forms
   a therapeutic composition is that comprises an anti-cancer drug selected from cytotoxic
   doxorubicin and taxol.
- 16. (Original) The method of claim 1, wherein the at least one encapsulant is encapsulated simultaneously with polymersome formation, or subsequent thereto.
- 17-22. (Cancelled)